Claims

1. A compound of formula (I) or a pharmaceutically or veterinarily acceptable salt. hydrate or solvate thereof

$$Q \xrightarrow{R_1} O \xrightarrow{R_4} N \xrightarrow{N} N \xrightarrow{N} A$$

$$Q \xrightarrow{R_2} R_3 \qquad (I)$$

wherein

Q represents a radical of formula -N(OH)CH(=O) or formula -C(=O)NH(OH);

Y represents -C(=O)-, -C(=S)-, -S(=O)-, or -SO₂-;

 R_1 represents hydrogen, C_1 - C_6 alkyl or C_1 - C_6 alkyl substituted by one or more halogen atoms, or, except when Q is a radical of formula -N(OH)CH(=O), a hydroxy, C_1 - C_6 alkoxy, C_1 - C_6 alkenyloxy, halogen, amino, C_1 - C_6 alkylamino, or di-(C_1 - C_6 alkyl)amino group;

 R_2 represents a substituted or unsubstituted C_1 - C_6 alkyl, C_1 - C_3 alkyl-O- C_1 - C_3 alkyl, C_1 - C_3 alkyl, cycloalkyl(C_1 - C_3 alkyl)-, aryl(C_1 - C_3 alkyl)-, heterocyclyl(C_1 - C_3 alkyl)-, or R^1R^2N - C_1 - C_3 alkyl group wherein R^1 represents hydrogen or C_1 - C_3 alkyl and R^2 represents C_1 - C_3 alkyl, or R^1R^2N - represents a cyclic amino group;

 R_3 and R_4 taken together with the nitrogen atoms to which they are respectively attached form a saturated heterocyclic ring of from 4 to 7 ring atoms, which may be fused to a second carbocyclic or heterocyclic ring, either of which rings may optionally be substituted; and

A represents a primary, secondary or tertiary amino group or a group - R_5 , - OR_5 , wherein R_5 is a substituted or unsubstituted C_1 - C_6 alkyl, C_2 - C_6 alkenyl, C_2 - C_6 alkynyl,

cycloalkyl, aryl, heterocyclyl, C_1 - C_3 alkyl-O- C_1 - C_3 alkyl, C_1 - C_3 alkyl-S- C_1 - C_3 alkyl, cycloalkyl(C_1 - C_3 alkyl)-, heterocyclic(C_1 - C_3 alkyl, aryl(C_1 - C_3 alkyl)-, or R^1R^2N - C_1 - C_3 alkyl group wherein R^1 represents hydrogen or C_1 - C_3 alkyl and R^2 represents C_1 - C_3 alkyl, or R^1R^2N - represents a cyclic amino group.

- 2. A compound as claimed in claim 1 wherein Q is an N-formyl hydroxylamine group -N(OH)CH(=O).
- 3 A compound as claimed in claim 1 or claim 2 wherein -Y- is -C(=O)- or SO₂.
- 4. A compound as claimed in any of the preceding claims wherein R_1 is hydrogen.
- 5. A compound as claimed in any of the preceding claims wherein R_2 is optionally substituted C_1 - C_6 alkyl, C_3 - C_6 alkenyl, C_3 - C_6 alkynyl or cycloalkyl; phenyl(C_1 - C_6 alkyl)-, phenyl(C_3 - C_6 alkenyl)- or phenyl(C_3 - C_6 alkynyl)- optionally substituted in the phenyl ring;

cycloalkyl(C_1 - C_6 alkyl)-, cycloalkyl(C_3 - C_6 alkenyl)- or cycloalkyl(C_3 - C_6 alkynyl)- optionally substituted in the cycloalkyl ring; or

 $CH_3(CH_2)_pO(CH_2)_{q^-}$ or $CH_3(CH_2)_pS(CH_2)_{q^-}$, wherein p is 0, 1, 2 or 3 and q is 1, 2 or 3.

6. A compound as claimed in any of claims 1 to 4 wherein R₂ is methyl, ethyl, nor iso-propyl, nor iso-butyl, nopentyl, iso-pentyl, 3-methyl-but-1-yl, nohexyl, nohexyl, noctyl, methylsulfanylethyl, ethylsulfanylmethyl, 2-methoxyethyl, 2-ethoxymethyl, 3-hydroxypropyl, allyl, 3-phenylprop-3-en-1-yl, prop-2-yn-1-yl, 3-phenylprop-2-yn-1-yl, 3-(2-chlorophenyl)prop-2-yn-1-yl, but-2-yn-1-yl, cyclopentyl, cyclopentylmethyl, cyclopentylpropyl,

acyclohexylmethyl, cyclohexylethyl, cyclohexylpropyl, furan-2-ylmethyl, furan-3-methyl, tetrahydrofuran-2-ylmethyl, tetrahydrofuran-2-ylmethyl, piperidinylmethyl, pyrid-2-ylmethyl, pyrid-3-ylmethyl, pyrid-4-ylmethyl, phenylpropyl, 4-chlorophenylpropyl, 4-methylphenylpropyl, 4-methoxyphenylpropyl, benzyl, 4-chlorobenzyl, 4-methylbenzyl, or 4-methoxybenzyl.

- 7. A compound as claimed in any of claims 1 to 4 wherein R_2 is (C_1-C_6) alkyl-, cycloalkylmethyl-, (C_1-C_3) alkyl-S- (C_1-C_3) alkyl-, or (C_1-C_3) alkyl-O- (C_1-C_3) alkyl-, especially n-propyl, n-butyl, n-pentyl, cyclopentylmethyl, cyclopentylethyl, cyclopentylmethyl or cyclohexylethyl.
- 8. A compound as claimed in any of the preceding claims wherein the ring formed by R_3 and R_4 and the nitrogens to which they are attached is one of the following, any of which may be optionally substituted, and wherein R represents hydrogen or C_1 - C_4 alkyl and any sulfur atom present as a ring member may be oxidised to -SO- or -SO₂-:

- 9. A compound as claimed in any of the preceding claims wherein A is a secondary amino group or a cyclic or non-cyclic tertiary amino group.
- 10. A compound as claimed in any of claims 1 to 8 wherein A is an amino group of formula NR_6R_7 wherein R_6 and R_7 independently represent a radical of formula (II)

$$\xrightarrow{} (Alk^1)_m - (X)_p - (Alk^2)_n - Z \qquad (II)$$

m, p and n are independently 0 or 1;

Z represents hydrogen or a carbocyclic or heterocyclic ring of 5 to 7 ring atoms which is optionally fused to a saturated or unsaturated carbocyclic or heterocyclic second ring of 5 to 7 ring atoms;

Alk¹ and Alk² independently represent divalent C₁-C₃ alkylene radicals;

X represents -O-, -S-, -S(O)-, -S(O₂)-, -C(=O)-, -NH-, -NR₇- where R₇ is C₁-C₃ alkyl;

and wherein

Alk¹, Alk² and Z when other than hydrogen, independently are optionally substituted by

(C₁-C₃)alkyl, (C₂-C₃)alkenyl, or (C₂-C₃)alkynyl, phenyl, optionally substituted by (C₁-C₃)alkyl, (C₁-C₃)alkoxy, halo, nitro, amino, mono- or di-(C₁-C₃)alkylamino, cyano or trifluoromethyl;

monocyclic 5 or 6-membered heterocyclic, optionally substituted by (C₁-C₃)alkyl, (C₁-C₃)alkoxy, halo, nitro, amino, mono- or di-(C₁-C₃)alkylamino, cyano or trifluoromethyl

benzyl, optionally substituted in the phenyl ring by (C_1-C_3) alkyl, (C_1-C_3) alkoxy, halo, nitro, amino, mono- or di- (C_1-C_3) alkylamino, cyano or trifluoromethyl,

hydroxy, phenoxy, (C_1-C_6) alkoxy, or hydroxy (C_1-C_6) alkyl, mercapto, (C_1-C_6) alkylthio or mercapto (C_1-C_6) alkyl,

охо,

nitro,

cyano

halo

- -COOH, or -COORA,
- -CONH₂, -CONHR^A, or -CONR^AR^B

-COR^A, -SO₂R^A,
-NHCOR^A,
-NH₂, -NHR^A, or -NR^AR^B,
wherein R^A and R^B are independently a (C₁-C₆) alkyl group, R^A and R^B taken together with the nitrogen atom to which they are attached form a 5- or 6-membered heterocyclic ring which may be substituted by (C₁-C₃)alkyl, hydroxy, or hydroxy(C₁-C₃)alkyl.

- 11. A compound as claimed in claim 10 wherein Alk¹ and Alk² independently represent -(CH₂)- or -(CH₂CH₂)-.
- 12. A compound as claimed in claim 10 or claim 11 wherein m is 0, p is 1, n is 0 or 1 and X is -C(=O)- or $-S(O_2)$ -.
- 13. A compound as claimed in claim 10 wherein the substituent (II) has the formula -CH₂Z, -OZ, or -(C=O)Z wherein Z is C₁-C₃ alkyl, phenyl, 3,4-methylenedioxyphenyl, morpholinyl, pyrimidinyl, 1,2,3-thiadiazolyl, 1,4-thiazolyl, benzofuranyl, furanyl, thienyl, pyranyl, pyrrolyl, pyrazolyl, isoxazolyl, or pyridyl, any of which may optionally be substituted as specified. In particular, Z may be a methyl, ethyl, n- or iso-propyl, phenyl, 3,4-methylenedioxyphenyl, morpholinyl, pyrimidin-2-yl, 1,2,3-thiadiazol-5-yl, 1,4-thiazol-5-yl, benzofuran-2-yl, 2-or 3-furanyl, 2- or 3-thienyl, 2- or 3-pyranyl, 2-, 3- or 4-pyrrolyl, 3-, 4- or 5-pyrazolyl, 3-, 4- or 5-isoxazolyl, or 2-, 3- or 4-pyridyl ring any of which may optionally be substituted as specified in the broad description of the compounds of the invention.
- 14. A compound as claimed in any of claims 1 to 8 wherein A is an amino group of formula -NR $_6$ R $_7$ wherein R $_6$ and R $_7$ taken together with the nitrogen atom to which they are attached form a saturated heterocyclic ring of 5 to 8 atoms optionally fused to a saturated or unsaturated carbocyclic or heterocyclic second ring of 5 to 7 ring atoms, any of which rings being optionally substituted by a radical of formula (II) as

defined in any of claims 10 to 13.

- 15. A compound as claimed in claim 14 wherein A is optionally substituted piperidin-1-yl or 1-piperazinyl.
- 16. A compound as claimed in claim 1 of formula (IIA) or (IIB)

wherein R₂ is as defined in claim 1;

X₁ is a bond, C₁-C₃ alkylene, -NH- or -O-; and

A₁ is optionally substituted C₁-C₆ alkyl, cycloalkyl, aryl, or heterocyclic.

17. A compound as claimed in claim 16 wherein R₂ is n-propyl, n-butyl, n-pentyl, cyclopentylmethyl, cyclopentylethyl, cyclohexylmethyl or cyclohexylethyl;

X₁ is a bond, -CH₂-, -CH₂CH₂-, -CH₂CH₂-, -NH- or -O-; and

A₁ is methyl, ethyl phenyl, cyclopentyl, cyclohexyl, 2- or 3-furanyl, 2- or 3- thienyl, 2-, 3- or 4-pyridyl, 3-, 4- or 5-pyrazolyl, 3-, 4- or 5-oxazolyl, or 3-, 4- or 5-thiazolyl, methoxymethyl, 3,5-bis-(trifluoromethyl)phenyl, 4-trifluoromethylphenyl, 4-methoxyphenyl, 3,4-methylenedioxyphenyl, 4-fluorophenyl benzyl, 3-pyridyl, 4-pyridyl, cyclohexyl, 1,3-dimethylpyrazol-5-yl, 1-methylimidazol-5-yl, or 2-[morpholin-1-yl]pyrid-5-yl.

18. A method for the treatment of bacterial infections in humans and non-human

WO 2004/050638 PCT/GB2003/005179

54

mammals, which comprises administering to a subject suffering such infection an antibacterially effective dose of a compound as claimed in any of the preceding claims.

- 19. The use of a compound as claimed in any of claims 1 to 17 for inhibiting bacterial growth in vitro and in vivo in mammals.
- 20. The use of a compound as claimed in any of claims 1 to 17 for the manufacture of a composition for treating bacterial infection by inhibiting bacterial growth.
- 21. A method for the treatment of bacterial contamination by applying an antibacterially effective amount of a compound as claimed in any of claims 1 to 17 to the site of contamination.
- 22. A pharmaceutical or veterinary composition comprising a compound as claimed in any of claims 1 to 17 together with a pharmaceutically or veterinarily acceptable carrier or excipient.